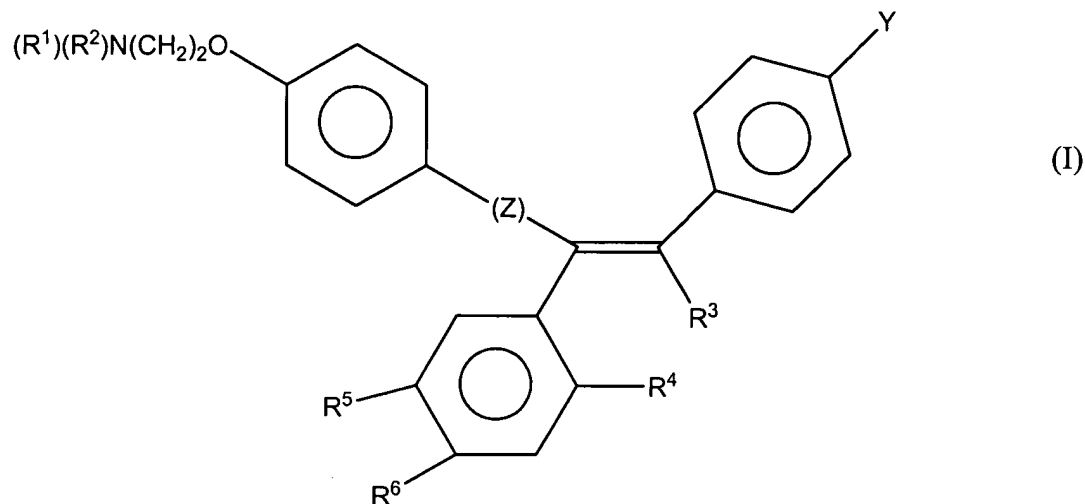


### IN THE CLAIMS

Please amend the claims as follows.

1-172. (Canceled).

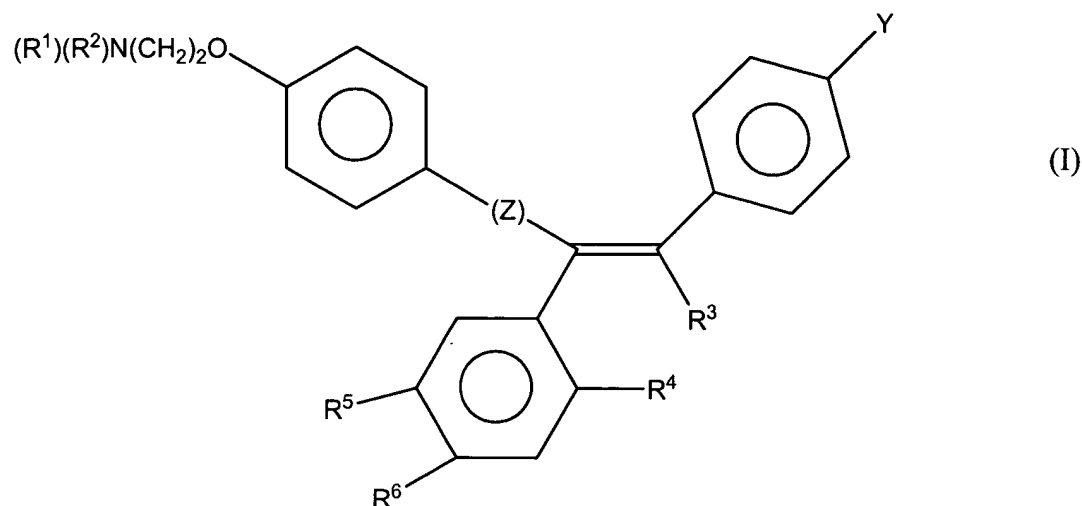
173. (Currently Amended) A therapeutic method for treating a cardiovascular or vascular indication characterized by a decreased lumen diameter comprising locally administering to a mammal at risk of or afflicted with said cardiovascular or vascular indication, a cytostatic dose of a therapeutic agent, wherein the therapeutic agent is a compound of formula (I):



wherein Z is C=O or a covalent bond; Y is H or O(C<sub>1</sub>-C<sub>4</sub>)alkyl, R<sup>1</sup> and R<sup>2</sup> are individually (C<sub>1</sub>-C<sub>4</sub>)alkyl or together with N are a saturated heterocyclic group, R<sup>3</sup> is ethyl or chloroethyl, R<sup>4</sup> is H, R<sup>5</sup> is I, O(C<sub>1</sub>-C<sub>4</sub>)alkyl or H and R<sup>6</sup> is I, O(C<sub>1</sub>-C<sub>4</sub>)alkyl or H with the proviso that when R<sup>4</sup>, R<sup>5</sup>, and R<sup>6</sup> are H, R<sup>3</sup> is not ethyl; or a pharmaceutically acceptable salt thereof.

174. (Previously Presented) The method of claim 173 wherein the cytostatic dose is effective to increase the level of TGF-beta so as to decrease lesion formation or development, decrease lipid accumulation, increase plaque stability, maintain or increase vessel lumen diameter, or any combination thereof.

175. (Previously Presented) The method of claim 173 wherein the compound of formula (I) is idoxifene, 4-iodotamoxifen, 3-iodotamoxifen, toremifene, or a pharmaceutically acceptable salt thereof.
176. (Previously Presented) The method of claim 173 wherein the compound of formula (I) is idoxifene or a pharmaceutically acceptable salt thereof.
177. (Previously Presented) The method of claim 173 wherein the compound of formula (I) is toremifene or a pharmaceutically acceptable salt thereof.
178. (Canceled).
179. (Previously Presented) The method of claim 173 wherein the compound of formula (I) is administered via a sustained release dosage form.
180. (Previously Presented) The method of claim 173 wherein the administration is localized at the site of vascular trauma.
181. (Previously Presented) The method of claim 173 wherein the compound directly or indirectly increases the level of active TGF-beta.
182. (Currently Amended) A therapeutic method of increasing the level of TGF-beta in a diabetic mammal at risk of or afflicted with a cardiovascular or vascular indication characterized by a decreased lumen vessel diameter, comprising administering to a diabetic mammal at risk of or afflicted with said cardiovascular or vascular indication an effective amount of a compound of formula (I):



wherein Z is C=O or a covalent bond; Y is H or O(C<sub>1</sub>-C<sub>4</sub>)alkyl, R<sup>1</sup> and R<sup>2</sup> are individually (C<sub>1</sub>-C<sub>4</sub>)alkyl or together with N are a saturated heterocyclic group, R<sup>3</sup> is ethyl or chloroethyl, R<sup>4</sup> is H or together with R<sup>3</sup> is -CH<sub>2</sub>-CH<sub>2</sub>- or -S-, R<sup>5</sup> is I, OH, O(C<sub>1</sub>-C<sub>4</sub>)alkyl or H and R<sup>6</sup> is I, O(C<sub>1</sub>-C<sub>4</sub>)alkyl or H with the proviso that when R<sup>4</sup>, R<sup>5</sup>, and R<sup>6</sup> are H, R<sup>3</sup> is not ethyl; or a pharmaceutically acceptable salt thereof.

183. (Previously Presented) The method of claim 182 wherein the increase in TGF-beta reduces diabetic retinopathy.

184. (Currently Amended) The method of claim 182 wherein the mammal is a human diabetic.

185. (Previously Presented) The method of claim 184 wherein the diabetic has retinopathy.

186. (Previously Presented) The method of claim 182 wherein the compound indirectly or directly increases the level of active TGF-beta in vascular tissue.

187. (Previously Presented) The method of claim 182 wherein the compound is a TGF-beta production stimulator.

188. (Previously Presented) The method of claim 182 wherein the compound is a TGF-beta activator.
189. (Previously Presented) The method of claim 182 wherein the compound increases the production of TGF-beta mRNA.
190. (Previously Presented) The method of claim 182 wherein the compound increases the cleavage of the latent form of TGF-beta.
191. (Previously Presented) The method of claim 182 wherein the compound increases the bioavailability of TGF-beta.
192. (Previously Presented) The method of claim 182 wherein the compound is idoxifene or a pharmaceutically acceptable salt thereof.
193. (Previously Presented) The method of claim 182 wherein the compound is toremifene or a pharmaceutically acceptable salt thereof.
194. (Previously Presented) The method of claim 182 wherein the compound is droloxifene or a pharmaceutically acceptable salt thereof.
195. (Canceled)
196. (Previously Presented) The method of claim 173 or 182 wherein the compound forms cellular DNA adducts at level which is reduced relative to DNA adduct formation by tamoxifen.
197. (Previously Presented) The method of claim 173 or 182 wherein the compound has estrogenic activity which is reduced relative to the estrogenic activity of tamoxifen.

198. (Previously Presented) The method of claim 173 or 182 wherein the compound does not form cellular DNA adducts.

199. (Previously Presented) The method of claim 173 or 182 wherein the compound has no estrogenic activity.

200. (Currently Amended) A method of increasing the level of TGF-beta in a mammal ~~at risk of or~~ afflicted with a cardiovascular ~~or vascular~~ indication characterized by a decreased lumen vessel diameter, comprising administering to a mammal ~~at risk of or~~ afflicted with said cardiovascular ~~or vascular~~ indication an effective amount of an agent that directly or indirectly elevates the level of active TGF-beta1 in said mammal, wherein the agent ~~has reduced estrogenic activity relative to tamoxifen, reduced DNA adduct formation relative to tamoxifen, or any combination thereof, and wherein the agent~~ is a structural analog of tamoxifen or a pharmaceutically acceptable salt thereof.

201. (Canceled).

202. (Previously Presented) The method of claim 200 wherein the agent is idoxifene or a pharmaceutically acceptable salt thereof.

203. (Previously Presented) The method of claim 200 wherein the agent is toremifene or a pharmaceutically acceptable salt thereof.

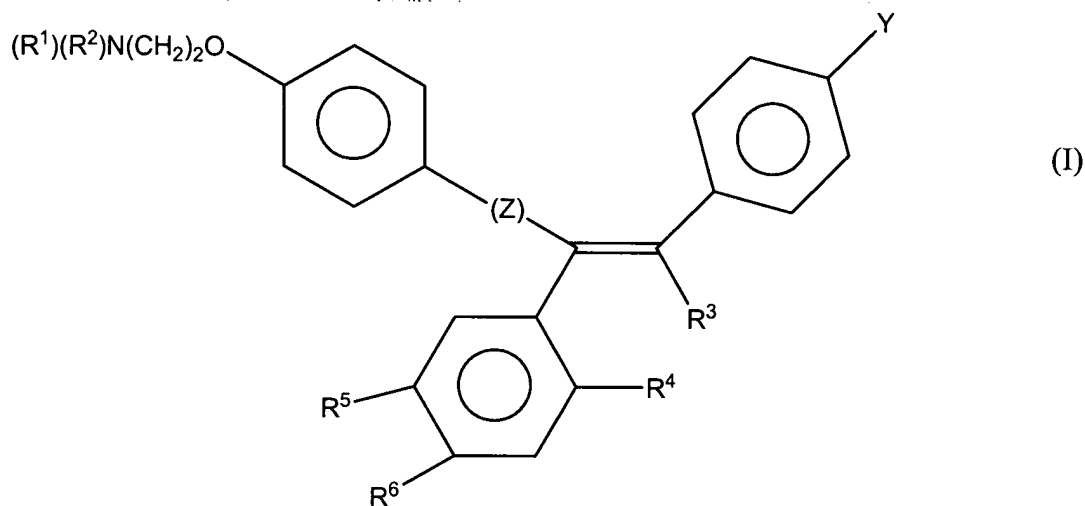
204. (Canceled).

205. (Previously Presented) The method of claim 173, 182, or 200 wherein the administration increases the level of latent TGF-beta relative to the level of latent TGF-beta prior to said administration.

206. (Previously Presented) The method of claim 173, 182, or 200 wherein the administration increases the level of active TGF-beta relative to the level of active TGF-beta prior to said administration.

207-230. (Canceled).

231. (Currently Amended) A therapeutic method for treating a condition selected from the group consisting of arteriosclerosis, silent myocardial infarction, vascular insufficiency in the limbs, peripheral neuropathy, and retinopathy and ~~small vessel disease~~, comprising administering to a mammal afflicted with said condition, an effective amount of a compound of formula (I):



wherein Z is C=O or a covalent bond; Y is H or  $O(C_1-C_4)alkyl$ ,  $R^1$  and  $R^2$  are individually  $(C_1-C_4)alkyl$  or together with N are a saturated heterocyclic group,  $R^3$  is ethyl or chloroethyl,  $R^4$  is H,  $R^5$  is I,  $O(C_1-C_4)alkyl$  or H and  $R^6$  is I,  $O(C_1C_4)alkyl$  or H with the proviso that when  $R^4$ ,  $R^5$ , and  $R^6$  are H,  $R^3$  is not ethyl; or a pharmaceutically acceptable salt thereof.

232-233. (Canceled).

234. (Currently Amended) The method of claim 173, 182, [[207]] or 231 wherein  $R^5$  or  $R^6$  is I.